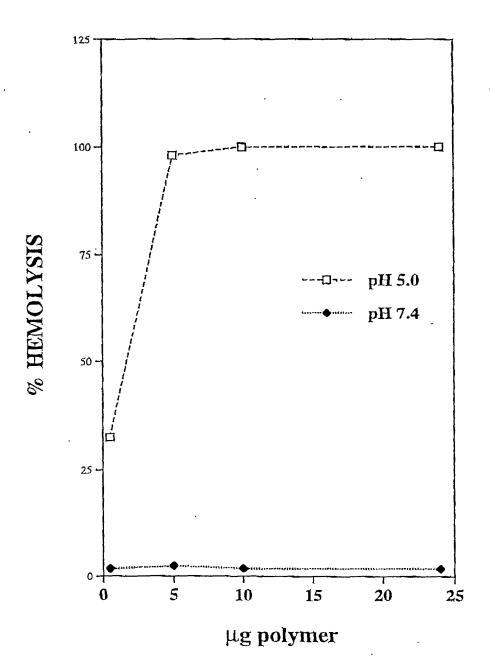
### HEMOLYSIS BY ACETAL-PEG-COPOLYMER



**Experimental Conditions** 

- (1) 2% RBCs in 1ml PBS buffer
- (2) Incubation temperature 37C
- (3) Incubation time 20 minutes
- (4) Experiments done in triplicate STD < 2%

## 

# Acid-Degradable Bonds Enhance Endosomal Drug Release of Targeted Polymer Carrier

endosome Disrupted Acid-degradable Endocytosis linker Membrane-disruptive polymer backbone "mask" backbone **Grafted PEGs Targeting** ligand

directly, or linked by PEG, -S-S- or (PEG/-S-S-) bonds DRUG ( ) may be conjugated, or complexed to backbone, each via an acid-degradable bond.

"Unmasked" backbone disrupts endosomal membrane

PEG-DRUG delivered into cytoplasm. Free DRUG or

4 Figure

SERUM STABLE NONTOXIC FORM MEMBRANE DISRUPTIVE FORM x = 50%y = 5%z = 45% Acid hydrolysis (Inside Endosome) pH degradable linkage PEG-Y X-PEG X-PEG

Encrypted Polymer E1: X = Y = Methoxy Encrypted Polymer E2: X = Fluorescein, Y = Lactose Encrypted Polymer E3: X = Hexalysine, Y = Lactose

Mercaptopropanol

## 

PTSOH in Dry THF
Basic Alumina
Chromotagraphy

Acetal Linkage

Hydroxypropyl-mercaptothiopyridal

Thiopyridal-amino-acetal Monomer

## Figure 5 Continued **AIBN** 60°C overnight 5kD Y-PEG-SH X-PEG-SH Ether Precipitation

Encrypted Polymer E1: X = Y = Methoxy Encrypted Polymer E2: X = Fluorescein, Y = Lactose Encrypted Polymer E3: X = Hexalysine, Y = Lactose

Y-PEG-S-S

